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(19) **United States**(12) **Patent Application Publication****Hinck et al.**(10) **Pub. No.: US 2021/0332096 A1**(43) **Pub. Date: Oct. 28, 2021**(54) **ENGINEERED TGF-BETA MONOMERS AND THEIR USE FOR INHIBITING TGF-BETA SIGNALING**

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**Publication Classification**(71) Applicants: **University of Pittsburgh - Of the Commonwealth System of Higher Education**, Pittsburgh, PA (US); **National Research Council of Canada**, Ottawa (CA)(51) **Int. Cl.**  
**C07K 14/495** (2006.01)(52) **U.S. Cl.**  
CPC ..... **C07K 14/495** (2013.01); **A61K 38/00** (2013.01)(72) Inventors: **Andrew Peterson Hinck**, Pittsburgh, PA (US); **Traian Sulea**, Ottawa (CA)(57) **ABSTRACT**(73) Assignees: **University of Pittsburgh - Of the Commonwealth System of Higher Education**, Pittsburgh, PA (US); **National Research Council of Canada**, Ottawa (CA)

Recombinant transforming growth factor (TGF)- $\beta$  monomers modified to inhibit dimerization and block TGF- $\beta$  signaling are described. The recombinant TGF- $\beta$  monomers lack the ability to bind and recruit TGF- $\beta$  type I receptor (T $\beta$ RI), but retain the capacity to bind the high affinity TGF- $\beta$  type II receptor (T $\beta$ RII), and in some instances, include mutations that increase their affinity for T $\beta$ RII. Nucleic acid molecules and vectors encoding the recombinant TGF- $\beta$  monomers are also described. Isolated cells, such as T cells, can be re-programmed with a TGF- $\beta$  monomer-encoding nucleic acid or vector to secrete the monomer. Use of the recombinant TGF- $\beta$  monomers and/or cells producing the recombinant TGF- $\beta$  monomers, to inhibit TGF- $\beta$  signaling, such as to treat disorders associated with aberrant TGF- $\beta$  signaling, are also described.

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**Specification includes a Sequence Listing.**